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IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

Applicant: Michalopoulos et al.

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For: A NOVEL LONG-TERM THREE DIMENSIONAL TISSUE CULTURE
SYSTEM

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INFORMATION DISCLOSURE STATEMENT

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42,328

PTO Registration No.

July 28, 2000

Date of Signature

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Assistant Commissioner of Patents
Washington, D.C. 20231

SIR:

In accordance with 37 C.F.R. § 1.56, Applicants respectfully request that the
references relating to the above-mentioned application listed herein in reverse chronological
alphabetical order be made of record in the U.S. Patent and Trademark Office.

1. Bonadio et al., August 24, 1999, U.S. Patent 5,942,496;
2. Caplan et al., January 5, 1999, U.S. Patent 5,855,619;
3. Elliot et al., June 8, 1999, U.S. Patent 5,910,582;
4. Michalopoulos, et al., 1999, "Morphogenetic events in mixed cultures of rat hepatocytes
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5. Mitaka et al., 1999, "Reconstruction of Hepatic Organoid by Rat Small Hepatocytes and Hepatic Nonparenchymal Cells", *Hepatology* 29:111-125;
 6. Mooney et al., March 23, 1999, U.S. Patent 5,885,829;
 7. Ni et al., June 8, 1999, U.S. Patent 5,910,431;
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 10. Cable, 1997, "Exposure of Primary Rat Hepatocytes in Long-Term DMSO Culture to Selected Transition Metals Induces Hepatocyte Proliferation and Formation of Duct-Like Structures", *Hepatology* 26:1444-1445;
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 19. Jukkola et al., 1993, "Procollagen synthesis and extracellular matrix deposition in MG-63

- osteosarcoma cells", *J. Bone Mineral Research* 8:651-657;
20. Li et al., December 14, 1993, U.S. Patent 5,270,192;
 21. Mikata et al., 1993, "Effects of Mitogens and Co-Mitogens on the Formation of Small-Cell Colonies in Primary Cultures of Rat Hepatocytes", *J. Cell Physiol.* 157:461-468;
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 23. Mikata et al., 1992, "Small Cell Colonies Appear in the Primary Culture of Adult Rat Hepatocytes in the Presence of Nicotinamide and Epidermal Growth Factor", *Hepatology* 16:440-447;
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 26. Kost et al., 1991, "Effect of 2% Dimethyl Sulfoxide on the Mitogenic Properties of Epidermal Growth Factor and Hepatocyte Growth Factor in Primary Hepatocyte Culture", *J. Cell Physiol.* 147:274-289;
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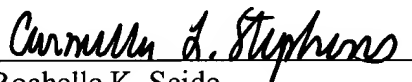
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35. Hagger et al., 1983, "Neonatal Hepatocyte Culture on Artificial Capillaries; A model for Drug Metabolism and the Artificial Liver", *ASAIO J.* 6:26-35;
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The referenced citations are listed in the accompanying PTO Form 1449. Copies of the references are submitted herewith.

Identification of the above-listed references is not to be construed as an admission of the Applicants or the attorneys of the Applicants that such references are available as "prior art" against the subject application.

Applicants respectfully request that the Examiner review the foregoing references and that the references be made of record in the file history of the above-mentioned application.

Respectfully submitted,



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